

(b) a recombinant complementing gene on an extrachromosomal vector, wherein the complementing gene is a functional replacement for said essential gene of (a) and wherein said complementing gene can recombine to replace the essential gene of (a); and

(c) a desired gene on the extrachromosomal vector, wherein the desired gene is a recombinant gene encoding a desired gene product;

wherein the desired gene is stably maintained in a progeny population of the microorganism.

46. (New) An attenuated derivative of a pathogenic microorganism which comprises:

(a) a non-functional native chromosomal essential gene;

(b) a recombinant complementing gene on an extrachromosomal vector, wherein the complementing gene can recombine to replace the non-functional chromosomal essential gene;

(c) a desired gene on the extrachromosomal vector, wherein the desired gene is a recombinant gene encoding a desired gene product; and

(d) an inactivating mutation in a native gene selected from the group consisting of a *pab* gene, a *pur* gene, and *aro* gene, *nadA*, *pncB*, *gale*, *pmi*, *fur*, *rpsL*, *ompR*, *htrA*, *hemaA*, *cdt*, *cya*, *crp*, *dam*, *phoP*, *phoQ*, *rfe*, *poxA*, *falU*, *mviA*, *sodC*, *recA*, *ssrA*, *sirA*, *inv*, *hila*, *rpoE*, *flgM*, *tonB*, and *slyA*;

wherein said complementing gene of (b) is a functional replacement for said essential gene of (a), wherein the desired gene is stably maintained in a progeny population of the microorganism.